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OPEN Clinical characteristics of external bacterial ocular and periocular infections and their antimicrobial treatment patterns among a Ghanaian ophthalmic population

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Empirical antimicrobial therapy is linked to a surge in antimicrobial resistant infections. However, an insight on the bacteria etiology of ocular infections is essential in the appropriation of choice of antimicrobial among clinicians, yet there remains a dearth of data from Ghana. We investigated the bacteria etiology of external ocular and periocular infections and antimicrobial treatment patterns among a Ghanaian ophthalmic population. A multicenter study design with purposive sampling approach was employed. Patients demographics and clinical data were collated using a pretested structure questionnaire. Cornea specimens and conjunctival swabs were obtained for bacterial isolation following standard protocols. About 95% (98/103) of ocular samples were positive for bacteria culture. The proportion of Gram-negative bacteria was 58.2%, and the predominant bacteria species were Pseudomonas aeruginosa 38.8% and Staphylococcus aureus 27.6%. Conjunctivitis 40.0% and keratitis 75.0% were mostly caused by Pseudomonas geruginosa. The routinely administered antimicrobial therapy were polymyxin B 41.2%, neomycin 35.1% and ciprofloxacin 31.6%. Participants demographic and clinical characteristics were unrelated with positive bacteria culture (p > 0.05). Our results showed a markedly high burden of ocular bacterial infections and variations in etiology. Bacterial infection-control and antimicrobial agent management programs should be urgently institutionalized to prevent the emergence of resistant infections.

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Abbreviations

Abbreviatio	118
A.E.H.	Anglican Eye Hospital
A.M.R.	Antimicrobial resistance
ARMOR	Antibiotic resistance monitoring in ocular microorganisms
C.H.A.G.	
C.H.R.P.E.	Committee on human research publication and ethics
C.I.P.	Ciprofloxacin
C.L.S.I.	Clinical laboratory standard institute guidelines
CONS	Coagulase negative Staphylococci species
C.O.R.	Crude odds ratio
D.N.A.	Deoxyribonucleic acid
FX	Flucoxacillin
KSH	Kumasi South Hospital
G.T.M.	Gentamycin
M.D.R.	Multi drug resistance
R.F.R.	Referral
PoB	Polymyxin B
NM	Neomycin
TBM	Tobramycin
TX	Tetracycline
O.X.T.	Oxytetracycline
OFC	Ofloxacin
R.N.A.	
S.M.H.	St Michaels Hospital

Globally, eye infections of bacterial origin remain a significant contributor to ocular morbidity and blindness, and the burden is increasing^{1,2}. Further, results from ocular microbial studies across different populations show no obvious pattern in prevalence estimates (ranging from 21.8 to 82.5%) across Africa²⁻¹², Asia¹³⁻²⁰, Australia^{21,22}, Europe²³⁻²⁵ and North America²⁶. Similarly, the bacterial etiology, thus the nature of Gram bacteria and the species of bacteria commonly implicated in external ocular (surface of the eyeball) and periocular (surrounding of the eyeball) infections vary across geographical regions and settings^{2-13,15-20,22,24-26}.

Anatomically, the eye is divided into three tunics: the outer (conjunctiva, sclera, cornea), middle (lens, ciliary body, iris), and inner (retina) coats. The tear film contains innate defenses such as bacteriocin, beta-lysin, lipocalin, lysozymes, immunoglobulins (Ig A, Ig G, Ig M), lactoferrins, and with antimicrobial effect against pathogenic strain of microorganisms. The external milieu of the eyes serves as a biome for pathogenic and non-pathogenic organisms. Naturally, antimicrobial constituents within the tear film prevent opportunistic microbes from causing any infections to the eyes. Dysregulation of the homeostatic balance as a result of trauma, contact lens wear, surgery, use of topical antibiotics, and reduced systemic immunity predisposes the eyes to opportunistic pathogens such as bacteria, fungi, virus, and protozoa. However, among these microbes, bacteria are commonly implicated in external ocular and periocular infections²⁷. Based on the exterior region affected; external ocular infections can be classified as blepharitis, conjunctivitis, keratitis, dacryocystitis, preseptal and orbital cellulitis^{28–30}, thus eyelids, conjunctiva, cornea, lacrimal sac, pre and post-septal areas are respectively involved.

These eye infections are usually treated with broad-spectrum antimicrobial agents with varying modes of action without any proper follow up for culture and sensitivity testing to identify the implicated pathogen³¹. In resource limited-environments such as Ghana, due to the unavailability of rapid diagnostic testing facilities, especially for identifying fastidious organisms, most clinicians: including eye care professionals engage in this prevailing empirical broad-spectrum antimicrobial treatment therapy³². Consequently, the irrational and prolonged use of broad-spectrum antibiotics in treating eye infections could alter the genetic makeup of ocular bacteria and consequently lead to antimicrobial resistance³³.

Ocular antimicrobial resistance is a growing public health threat in both advanced and developing countries. In the developed world nationwide surveillance programs have been institutionalized to monitor ascendancies in the antimicrobial resistance curve and subsequently tackle it. Among these nationwide surveillance programs include Ocular Tracking Resistance in the U.S. Today (Ocular TRUST)³⁴, Antibiotic Resistance Monitoring in Ocular Microorganisms (ARMOR)³⁵, European Antimicrobial Resistance Surveillance System³⁶, and Swedish Strategic Programme for the Rational Use of Antimicrobial Agents and Surveillance of Resistance³⁷. However, in developing countries such as Ghana, aside from the limited accessibility to a national antimicrobial policy, there are compromise regulatory measures and poor adherence to the use of antibiotics³⁸.

Multidrug resistance (MDR) is gradually gaining attention in mainstream medicine and public healthcare generally, as it renders antimicrobial agents inefficacious against pathogenic strains of bacteria. MDR causes delays in treatment and recoveries, rise in cost of therapy as well as increase in hospitalization time³⁹⁻⁴¹. In a nationwide laboratory based surveillance studies in Ghana, Opintan et al. reported over 70% prevalence of MDR among antibiotics such chloramphenicol, gentamycin, tetracycline, and quinolones against isolated bacterial strains from various infections of the urine, blood, sputum, ears, and eyes⁴². This finding was consistent with studies conducted in the People's Republic of China^{43,44}, Italy⁴⁵, and Ethiopia¹² which showed similar increasing trends. Consequentially, without drastic measures, it is estimated that the world will experience over 10 million annual AMR-related deaths hence it has become imperative to devise ammunitions to curb the situation⁴⁶. Importantly, a decline in ocular MDR will result in increased life expectancy as ocular resistance infections and associated blindness induce mortality.

There is a paucity of data on the prevalence and bacteria etiology of ocular infections in Ghana^{9,27}. Furthermore, earlier microbiological investigations in Ghana did not exquisitely focus on ocular infections and associated microbes^{42,47–49}. The absence of country-specific contemporary estimates limit the modeling of future scenarios, and assumptions with unreliable data and/or making decisions with evidence from other countries is of questionable utility given the geographic differences. An insight on bacteria etiology of ocular infections presented by Ghanaian patients is critical for desirable choice of antibiotic therapy by clinicians. Therefore, the study aims to investigate the bacteria etiology of external ocular and periocular infections, and antimicrobial treatment patterns among a Ghanaian ophthalmic population. The isolates recovered from ocular specimen will aid future antibiotic sensitivity studies and also serve as a gateway for exploration of local medicinal plants as alternative therapeutic agents.

Results

Description of the sample. Table 1 presents the sociodemographic, socioeconomic and healthcare status characteristics of the study participants. Out of the 114 patients presenting with external ocular and periocular infections, majority were females (56.1%), of median age of 17.0 (Interquartile range; 29.75) years (Table 1). The majority of the participants were aged 3–17 years (29.8%), of Akan ethnicity (93.0%), and with a protestant religion (80.7%). Most of them lived in a rural community (59.6%), with their highest education level being primary (33.6%) and major occupation as students (43.9%). An equal proportion were single (21.2%) and married (21.2%) and the remaining either cohabiting (3.5%), divorced (2.7%), widow (1.8%) or separated (0.9%). A preponderance of participants never smoked (52.6%), and with a significantly higher average alcohol intake in males compared to females (p=0.027). Approximately 6% had hypertension and an equal proportion had diabetes (4.4%) and peptic ulcer (4.4%). The number of patients on antihypertensive, antidiabetic and antibiotic medications were 5.3%, 3.5% and 3.5%, respectively.

Clinical characteristics of external ocular and periocular infections among study partici-Table 2 shows the clinical characteristics of external ocular and periocular infections among study pants. participants. Most presented with both eyes infected (66.7%) with the conjunctiva (94.7%) as the commonest affected site. The majority of the participants had previously used antimicrobials (55.8%) and have had previous eye infections (45.1%), and with duration of infection less than one-week (63.7%). There was a significant variation between males and females in terms of previous ocular trauma (p = 0.023) and previous use of mascara (p=0.015). With respect to monocular visual acuity, majority had right eye visual acuity better than 6/18(63.8%) and with a fewer having visual acuity worse than 6/60 (2.6%). Most had a left visual acuity within the ranges of 6/5-6/6 (67.5%) and fewer proportion (1.8%) having visual acuity better than 6/60 but worse than 6/24. The commonest presenting symptom was hyperemia/redness (69.3%), followed by discharge, (62.3%) itching (60.5%), eye pain (50.9%) and a smaller fraction having falling lashes (0.9%). Majority received a minimum of two antibiotics for treatment of infections (40.4%) and with a fewer portion (7.9%) having treatment other than antibiotics. The proportion of clinical presentations were conjunctivitis (60.5%), keratoconjunctivitis (11.4%), blepharoconjunctivis (9.6%), keratitis (7.9%), ocular trauma (6.1%), hordeolum (2.6%), preseptal cellulitis (0.9%) and ophthalmia neonatorum (0.9%). Other associated conditions were dry eyes (10.5%), headaches (5.3%), pterygium (4.4%) and pinguecula (0.9%). The commonly used antimicrobial therapeutics were polymyxin B (41.2%), neomycin (35.1%) and ciprofloxacin (31.6%) and fewer instances gentamycin (2.6%) and ofloxacin (1.6%) as shown in Table 3.

Bacteria etiology of external ocular and periocular infections among study participants. One hundred and three (103/114) ocular specimens were enrolled for bacterial isolation owing to the inability to obtain swabs from uncooperative minor subjects. Ninety-eight (95.1%) of the samples were culture positive, and no mixed cultured was identified (Please see Table 3). The proportion of Gram-negative bacteria was 58.2% with *Pseudomonas aeruginosa* (38.8%) and *Pseudomonas putida* (11.2%) being the predominant species. Conversely, the prevalence of Gram-positive bacteria was 41.8%, with a preponderance of bacteria pathogens being *Staphylococcus aureus* (27.6%) and *Coagulase negative staphylococci, CONS* (13.3%). The commonest strains of bacteria pathogens isolated from conjunctivitis were *Pseudomonas aeruginosa* (40.0%), *Staphylococcus aureus* (21.8%) and *Pseudomonas putida* (16.4%). Similarly, *Pseudomonas aeruginosa* (41.7%) and *Staphylococcus aureus* (33.3%) were frequently identified in cases of keratoconjunctivitis. *Staphylococcus aureus* (100.0%) was the sole organism implicated in cases of preseptal cellulitis, whereas most cases of keratitis was caused by *Pseudomonas aeruginosa* (75.0%) as shown in Table 4.

Factors associated with external ocular and periocular infections. Table 5 shows logistic regression analyses of the association between patients' demographics, clinical characteristics and prevalence of bacterial infection. None of the factors was significantly associated with prevalence of bacterial ocular infections (p > 0.05).

The novel study for the first time aims to investigate the bacteria etiology of external ocular and periocular infections and antimicrobial treatment patterns among a Ghanaian ophthalmic population. About 95% of the culture were positive for bacteria pathogens, and with the predominant class of bacteria being Gram negatives. *Pseudomonas aeruginosa* and *Staphylococcus aureus* were the commonly isolated bacteria species and with the former frequently isolated in cases of conjunctivitis and keratitis. The commonly used antimicrobial therapy in the clinical management of eye infections in these facilities were polymyxin B, neomycin and ciprofloxacin.

Bacteria ocular and periocular infections pose health challenges owing to associated morbidity and blindness. Globally, the burden of bacteria eye infections is higher especially in lower-and-middle income countries

	Total (N=114)	Males (N = 50, 43.9%)	Females (N = 64, 56.1%)	_
Variables	% (frequency)	% (frequency)	% (frequency)	<i>p</i> -value for linearity
Demographic characteristics				
Age group (years)				
0-2	21.1 (24)	20.0 (10)	21.9 (14)	0.948
3-7	29.8 (34)	32.0 (16)	28.1 (18)	
18-39	27.2 (31)	28.0 (14)	26.6 (17)	
≥40	21.9 (25)	20.0 (10)	23.4 (15)	
Ethnicity [‡]				
Akan	93.0 (106)	90.0 (45)	95.3 (61)	0.271
Northerner	7.0 (8)	10.0 (5)	4.7 (3)	
Religion				
Catholic	13.2 (15)	22.0 (11)	6.3 (4)	0.041
Protestant	80.7 (92)	760 (38)	84.4 (54)	
Muslim	4.4 (5)	2.0 (1)	6.3 (4)	
Atheist	1.8 (2)	0.0 (0)	3.1 (2)	
Socioeconomic characteristics	I			
Residence				
Rural	59.6 (68)	60.0 (30)	59.4 (38)	0.946
Urban	40.4 (46)	40.0 (20)	40.6 (26)	
Highest level of education	1	1	1	1
None	4.4 (5)	4.0 (2)	4.8 (3)	0.943
Preschool	13.3 (15)	12.0 (6)	14.3 (9)	
Primary	33.6 (38)	30.0 (15)	36.5 (23)	
Secondary	29.2 (33)	32.0 (16)	27.0 (17)	
Tertiary	8.0 (9)	10.0 (5)	6.3 (4)	
Not applicable*	11.5 (13)	12.0 (6)	11.1 (7)	
Occupation				
Farming	4.4 (5)	6.0 (3)	3.1 (2)	0.092
Wage/Salary worker	7.0 (8)	6.0 (3)	7.8 (5)	
Construction worker	2.6 (3)	6.0 (3)	0.0 (0)	
Dressmaking	1.8 (2)	0.0 (0)	3.1 (2)	
Driver/Transport business	2.6 (3)	6.0 (3)	0.0 (0)	
Businessman/woman	5.3 (6)	4.0 (2)	6.3 (4)	
Welding	1.8 (2)	4.0 (2)	0.0 (0)	
Trading	8.8 (10)	4.0 (2)	12.5 (8)	
Beautician	1.8 (2)	0.0 (0)	3.1 (2)	
Hairdressing	2.6 (3)	2.0 (1)	3.1 (2)	
Student	43.9 (50)			
Unemployed	5.3 (6)	46.0 (23) 2.0 (1)	42.2 (27) 7.8 (5)	
		14.0 (7)		
Not applicable*	12.3 (14)	14.0 (7)	10.9 (7)	
Marital status	40.7 (55)	50.0 (25)	47 ((20)	0.026
Not applicable*	48.7 (55)	50.0 (25)	47.6 (30)	0.936
Single	21.2 (24)	24.0 (12)	19.0 (12)	
Married	21.2 (24)	18.0 (9)	23.8 (15)	
Cohabiting	3.5 (4)	4.0 (2)	3.2 (2)	
Divorced	2.7 (3)	2.0 (1)	3.2 (2)	
Separated	0.9 (1)	0.0 (0)	1.6 (1)	
Widow	1.8 (2)	2.0 (1)	1.6 (1)	
Health status variables				
Smoking habits				1
Not applicable*	44.7 (51)	46.0 (23)	43.8 (28)	0.237
Never smoked	52.6 (60)	48.0 (24)	56.3 (36)	
Past smoker	1.8 (2)	4.0 (2)	0.0 (0)	
Current smoker	0.9 (1)	2.0 (1)	0.0 (0)	
Alcohol intake habits		1	1	1
Not applicable*	43.9 (50)	42.0 (21)	45.3 (29)	0.054

	Total (N=114)	Males (N = 50, 43.9%)	Females (N=64, 56.1%)		
Variables	% (frequency)	% (frequency)	% (frequency)	<i>p</i> -value for linearity	
I never drink	46.5 (53)	40.0 (20)	51.6 (33)		
I drink only on special occasions	7.9 (9)	14.0 (7)	3.1 (2)		
I drink once or twice a week	1.8 (2)	4.0 (2)	0.0 (0)		
Average alcohol consumption per	week				
None	90.4 (103)	82.0 (41)	96.9 (62)	0.027	
1 unit	6.1 (7)	12.0 (6)	1.6 (1)		
2–5 unit	3.5 (4)	6.0 (3)	1.6 (1)		
Medical history					
Diabetes	4.4 (5)	2.0 (1)	6.3 (4)	0.272	
Hypertension	6.1 (7)	4.0 (2)	7.8 (5)	0.4	
Tuberculosis	0.9 (1)	0.0 (0)	1.6 (1)	0.375	
Sexually transmitted diseases	0.9 (1)	2.0 (1)	0.0 (0)	0.256	
Peptic ulcer	4.4 (5)	2.0 (1)	6.3 (4)	0.272	
Others	2.6 (3)	6.0 (3)	0.0 (0)	0.047	
Is condition ongoing					
Not applicable*	80.7 (92)	82.0 (41)	79.7 (51)	0.915	
No	2.6 (3)	2.0 (1)	3.1 (20		
Yes	16.7 (19)	16.0 (8)	17.2 (11)		
Are you currently taking any med	ication				
No	82.5 (94)	82.0 (41)	82.8 (53)	0.91	
Yes	17.5 (20)	18.0 (9)	17.2 (11)		
Class of medications			1		
Antibiotic	3.5 (4)	6.0 (3)	1.6 (1)	0.201	
Antidiabetic	3.5 (4)	2.0 (1)	4.7 (3)	0.439	
Antihypertensive	5.3 (6)	4.0 (2)	6.3 (4)	0.593	
Antimalarial	0.9 (1)	0.0 (0)	1.6 (1)	0.375	
Anticholesterol	0.9 (1)	2.0 (1)	0.0 (0)	0.256	
Others	7.9 (9)	8.0 (4)	7.8 (5)	0.971	

 Table 1. Description of the sample. *Under age; %, percentage frequency. *Northerner is a collective name for all ethnic group in the northern region of Ghana.

including Ghana²⁻⁴. Though microscopic, the wide biodiversity of bacteria pathogens makes it burdensome for ophthalmic clinicians and physicians when selecting appropriate antibiotic therapy in routine clinical management of ocular and periocular infections. Previously, authors from several geographical jurisdictions have investigated the burden and etiology of bacteria eye infections, however, outcomes from these studies varied considerably^{13-15,18,20,25,26}. The prevalence estimates of medical conditions such as bacteria ocular and periocular infections are critical in informing eye care service delivery and in the development of policies to strengthen eye care practices yet there is presently limited ophthalmic data to propagate such transitions within the Ghanaian context. Importantly, gaining insight on bacteria etiology implicated in cases of external ocular and periocular infections within the Ghanaian population is essential to guide clinicians in the appropriate choice of antimic crobial therapy. Nonetheless there is paucity of data in this regard.

Overall, the prevalence of bacteria ocular and periocular infections found in this study was 95.1%. Our results are comparable with studies in Ethiopia⁵⁰, Saudi Arabia⁵¹, Italy²⁵ and United States of America²⁶. In a cross-sectional study in Ethiopia, Tesfaye et al. reported a prevalence of 74%⁵⁰. Similarly, a study by Shahaby and colleagues utilizing participants from a university clinic in Saudi-Arabia found more than two thirds of ocular specimens harboring bacteria pathogens⁵¹. Likewise, an observational case series conducted in Italy by Papa and coworkers, revealed that the proportion of bacterial infections was estimated at 72.5%²⁵. Furthermore, in a prospective observational study among patients undergoing cataract surgery in the U.S.A., Ta et al. showed that almost eight of every ten ocular specimen obtained from patients eyes had a bacteria etiology²⁶. On the one hand, estimates from the present study is significantly higher and varies substantially compared to studies in China^{13,18}, Iran²⁰, South Korea¹⁴ and Nepal¹⁵ with prevalence estimates far lower than 50%^{13–15,18,20}. Although geographical settings, study population, seasonality and laboratory procedures could account for such variations as reported earlier, a plausible reason for our observation may be attributable to the fact that our study unlike previous investigations enrolled patients from multiple eye care facilities hence the burden of infections maybe summative. Another reason is that majority of our study participants were rural dwellers with sanitation in such areas usually problematic compared to inhabitants in urban vicinities.

We observed a slightly higher proportion of Gram-negative bacteria compared to Gram positives as etiological agent in our study. This findings contrast with studies from China¹³, Ethiopia¹¹, Saudi-Arabia¹⁶, Uganda⁴ and

	Total (N=114)	Males (N = 50, 43.9%)	Females (N = 64, 56.1%)	
	% (frequency)	% (frequency)	% (frequency)	<i>p</i> -value for linearity
Eyes affected		1		
Right eye; oculus dexter	16.7 (19)	22.0 (11)	12.5 (8)	0.369
Left eye; oculus sinister	16.7 (19)	14.0 (7)	18.8 (12)	
Both eyes; oculus uterque	66.7 (76)	64.0 (32)	68.8 (44)	
Site of eye affected				
Eyelid/eye lashes	20.2 (23)	22.0 (11)	18.8 (12)	0.668
Conjunctiva	94.7 (108)	92.0 (46)	96.9 (62)	0.247
Cornea	18.4 (21)	26.0 (13)	12.5 (8)	0.065
Risk factors		·		
Previous eye infections	45.1 (51)	44.0 (22)	46.0 (29)	0.829
Previous use of antimicrobials	55.8 (63)	52.0 (26)	58.7 (37)	0.474
Previous usage of contact lenses	0.9 (1)	0.0 (0)	1.6 (1)	0.371
Previous use of spectacles	9.7 (11)	10.0 (5)	9.5 (6)	0.932
Previous ocular trauma	20.4 (23)	30.0 (15)	12.7 (8)	0.023
Previous ocular surgery	2.7 (3)	2.0 (1)	3.2 (2)	0.700
Duration of illness	1	1		
<1 week	63.7 (72)	66.0 (33)	61.9 (39)	0.649
2–4 weeks	18.6 (21)	20.0 (10)	17.5 (11)	
>4 weeks	17.7 (20)	14.0 (7)	20.6 (13)	
Previous application of mascara	6.2 (7)	0.0 (0)	11.1 (7)	0.015
Previous application of breastmilk	2.7 (3)	2.0 (1)	3.2 (2)	0.700
Presenting visual acuity		.,,		
Right eye; oculus dexter				_
Unavailable	2.6 (3)	2.0 (1)	3.1 (2)	0.939
FFL	23.7 (27)	24.0 (12)	23.4 (15)	
6/5-6/18	63.2 (72)	62.0 (31)	64.1 (41)	
6/24-6/60	7.9 (9)	8.0 (4)	7.8 (5)	
3/60-1/60	2.6 (3)	4.0 (2)	1.6 (1)	
Left eye; oculus dexter	2.0 (0)	1.0 (2)	110 (1)	
Unavailable	2.6 (3)	2.0 (1)	3.1 (2)	0.356
FFL	23.7 (27)	24.0 (12)	23.4 (15)	0.550
6/5-6/18	67.5 (77)	66.0 (33)	68.8 (44)	
6/24-6/60	1.8 (2)	0.0 (0)	3.1 (2)	
3/60-1/60	4.4 (5)	8.0 (4)	1.6 (1)	
Both eyes; oculus uterque	4.4 (3)	8.0 (4)	1.0 (1)	
Unavailable	91.2 (104)	86.0 (43)	95.3 (61)	0.266
FFL	0.9 (1)	2.0 (1)	0.0 (0)	0.200
	-			
6/5-6/18	7.0 (8)	10.0 (5)	4.7 (3)	
3/60-1/60	0.9 (1)	2.0 (1)	0.0 (0)	
Presenting patient symptoms	50.0 (59)	42.0 (21)	57 8 (27)	0.004
Eye pain Itabing	50.9 (58)	42.0 (21)	57.8 (37)	0.094
Itching Falling lashes	60.5 (69)	54.0(27)	65.6 (42)	0.208
Falling lashes	0.9 (1)	0.0 (0)	1.6 (1)	0.375
Lacrimation/watering	61.4 (70)	64.0 (32)	59.4 (38)	0.615
Hyperemia/redness	69.3 (79)	72.0 (36)	67.2 (43)	0.580
Swelling	21.9 (25)	22.0 (11)	21.9 (14)	0.987
Discharge	62.3 (71)	70.0 (35)	56.3 (33)	0.133
Burning sensation	7.9 (9)	6.0 (3)	9.4 (6)	0.507
Foreign body sensation	10.5 (12)	14.0 (7)	7.8 (5)	0.285
Others	11.4 (13)	4.0 (2)	17.2 (11)	0.028
Test/investigations				1
Visual acuity	99.1 (113)	98.0 (49)	100 (64)	0.256
Slit lamp Biomicroscopy	99.1 (113)	98.0 (49)	100 (64)	0.256
Ophthalmoscopy	98.2 (112)	98.0 (49)	98.4 (63)	0.860
Microbial analysis	90.4 (103)	88.0 (44)	92.2 (59)	0.452

	Total (N=114)	Males (N = 50, 43.9%)	Females (N=64, 56.1%)	
	% (frequency)	% (frequency)	% (frequency)	p-value for linearity
Clinical signs				
Eyelashes				
Healthy	98.2 (112)	100.0 (50)	96.9 (62)	0.451
Misdirected	0.9 (1)	0.0 (0)	1.6 (1)	
Crust formation	0.9 (1)	0.0 (0)	1.6 (1)	
Eyelids				
Healthy	66.7 (76)	66.0 (33)	67.2 (43)	0.819
Papillae	17.5 (20)	18.0 (9)	17.2 (11)	
Cobblestones	1.8 (2)	2.0 (1)	1.6 (1)	
Swelling	11.4 (13)	12.0 (6)	10.9 (7)	
Crust	0.9 (1)	2.0 (1)	0.0 (0)	
Drooping	0.9 (1)	0.0 (0)	1.6 (1)	
Rashes	0.9 (1)	0.0 (0)	1.6 (1)	
Conjunctiva	I			
Healthy	26.3 (30)	28.0 (14)	25.0 (16)	0.646
Injection	71.9 (82)	72.0 (36)	71.9 (46)	
Limbal papillae	0.9 (1)	0.0 (0)	1.6 (1)	
Limbal pigmentation	0.9 (1)	0.0 (0)	1.6 (1)	
Cornea	I	1	<u>.</u>	1
Transparent	87.7 (100)	78.0 (39)	95.3 (61)	0.045
Opacities	3.5 (4)	4.0 (2)	3.1 (2)	
Laceration	0.9 (1)	2.0 (1)	0.0 (0)	
Abrasions	7.0 (8)	14.0 (7)	1.6 (1)	
Ulcer	0.9 (1)	2.0 (1)	0.0 (0)	
Sclera		.,,		
Healthy	83.3 (95)	78.0 (39)	87.5 (56)	0.177
Pigmented	16.7 (19)	22.0 (11)	12.5 (8)	
Anterior chamber				
Deep	98.2 (112)	96.0 (48)	100 (64)	0.106
Shallow	1.8 (2)	4.0 (2)	0.0 (0)	
Pupils				
Healthy (PERLLA)	99.1 (113)	98.0 (49)	100.0 (64)	0.256
Abnormal (RAPD)	0.9 (1)	2.0 (1)	0.0 (0)	
Iris	015 (1)	210 (1)	0.0 (0)	
Health (Dark and flat)	99.1 (113)	98.0 (49)	100.0 (64)	0.256
Prolapse	0.9 (1)	2.0 (1)	0.0 (0)	0.230
Lens	0.5 (1)	2.0 (1)	0.0 (0)	
Transparent	94.7 (108)	92.0 (46)	96.9 (62)	0.247
Opacities	5.3 (6)	8.0 (4)	3.1 (2)	0.247
Number of antibiotics adminis		0.0 (4)	5.1 (2)	
None	7.9 (9)	6.0 (3)	9.4 (6)	0.351
				0.331
1 2	33.3 (38)	38.0 (19)	29.7 (19)	
	40.4 (44)	44.0 (22)	37.5 (24)	
≥3	18.4 (21)	12.0 (6)	23.4 (15)	
Clinical presentation	(0.5.(0))	52.0 (20)	(7.2.(42))	0.216
Conjunctivitis	60.5 (69)	52.0 (26)	67.2 (43)	0.216
Blepharoconjuntivitis	9.6 (11)	10.0 (5)	9.4 (6)	
Keratoconjunctivitis	11.4 (13)	16.0 (8)	7.8 (5)	
Ophthalmia neonatorum	0.9 (1)	2.0 (1)	0.0 (0)	
Ocular trauma	6.1 (7)	8.0 (4)	4.7 (3)	
Preseptal cellulitis	0.9 (1)	0.0 (0)	1.6 (1)	
Hordeolum	2.6 (3)	0 (0.0)	3 (4.7)	
Keratitis	7.9 (9)	6 (12.0)	3 (4.7)	
Associated conditions				
Dry eye syndrome		6.0 (3)		

	Total (N = 114)	Males (N = 50, 43.9%)	Females (N=64, 56.1%)	
	% (frequency)	% (frequency)	% (frequency)	<i>p</i> -value for linearity
Pterygium	4.4 (5)	4.0 (2)	4.7 (3)	0.859
Pingueculum	0.9 (1)	0.0 (0)	1.6 (1)	0.375
Headaches	5.3 (6)	8.0 (4)	3.1 (2)	0.247

Table 2. Clinical characteristics of external ocular and periocular infections among a Ghanaian ophthalmic population. *FFL* fixate-and-follow light, *PERLLA* pupils are equal, round, and reactive to light and accommodation, *RAPD* relative afferent pupillary defect.

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	Bacteria cul n (%)	ture results	ts Antimicrobial therapy employed in treating external ocular and periocular infections n (%)									
Types of clinical presentation	Positive	Negative	RFR	CIP	GTM	РоВ	NM	ТВМ	ТХ	OTX	OFC	FX
Conjunctivitis	55 (94.8)	3 (5.2)	0 (0.0)	22 (31.9)	2 (2.9)	29 (42.0)	22 (31.9)	13 (18.8)	16 (23.2)	12 (2.6)	0 (0.0)	0 (0.0)
Blepharoconjuntivitis	11 (100.0)	0 (0.0)	0 (0.0)	2 (18.2)	0 (0.0)	8 (72.7)	5 (45.5)	1 (9.1)	2 (18.2)	3 (27.3)	0 (0.0)	0 (0.0)
Keratoconjunctivitis	12 (92.3)	1 (7.7)	0 (0.0)	2 (15.4)	0 (0.0)	5 (38.5)	9 (69.2)	0 (0.0)	4 (30.8)	0 (0.0)	0 (0.0)	0 (0.0)
Ophthalmia neonatorum	1 (100.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (100.0)	0 (0.0)	1 (100.0)	0 (0.0)	1 (100.0)	0 (0.0)	0 (0.0)
Ocular trauma	7 (100.0)	0 (0.0)	1 (14.3)	3 (42.9)	0 (0.0)	1 (14.3)	1 (14.3)	0 (0.0)	2 (28.6)	0 (0.0)	0 (0.0)	1 (14.3)
Preseptal cellulitis	1 (100.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (100.0)	0 (0.0)	1 (100.0)	0 (0.0)	0 (0.0)	1 (100.0)
Hordeolum	3 (100.0)	0 (0.0)	0 (0.0)	1 (33.3)	0 (0.0)	1 (33.3)	1 (33.3)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	3 (100.0)
Keratitis	8 (88.9)	1 (11.1)	0 (0.0)	6 (66.7)	1 (11.1)	2 (22.2)	1 (11.1)	1 (11.1)	3 (33.3)	2 (22.2)	1 (11.1)	0 (0.0)
Total	98 (95.1)	5 (4.4)	1 (0.9)	36 (31.6)	3 (2.6)	47 (41.2)	40 (35.1)	16 (14.0)	28 (24.6)	18 (15.8)	1 (0.9)	5 (4.4)

Table 3. Antimicrobial treatment of external ocular and periocular infections in a Ghanaian ophthalmic population. *n* frequency, % percentage frequency, *RFR* referral, *CIP* ciprofloxacin, *GTM* gentamycin, *PoB* polymyxin B, *NM* neomycin, *TBM* tobramycin, *TX* tetracycline, *OXT* oxytetracycline, *OFC* ofloxacin, *FX* flucoxacillin.

	Types of clinical presentation								
Bacteria isolates	Conjunctivitis n (%)	Blepharoconjuntivitis n (%)	Keratoconjunctivitis n (%)	Ophthalmia neonatorum n (%)	Ocular trauma n (%)	Preseptal cellulitis n (%)	Hordeolum n (%)	Keratitis n (%)	Total isolates n (%)
Gram positive							•		
S. aureus	12 (21.8)	8 (72.7)	4 (33.3)	0 (0.0)	1 (14.3)	1 (100.0)	0 (0.0)	1 (12.5)	27 (27.6)
CONS	8 (14.5)	2 (18.2)	1 (8.3)	1 (100.0)	1 (14.3)	0 (0.0)	0 (0.0)	0 (0.0)	13 (13.3)
Streptococcus spp	1 (1.8)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (1.0)
Gram negative		1			1				-
Citrobacter spp	1 (1.8)	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	0 (0.0)	0 (0.0)	0 (0.0)	2 (2.0)
Serratia spp	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (33.3)	0 (0.0)	1 (1.0)
Salmonella spp	1 (1.8)	0 (0.0)	1 (8.3)	0 (0.0)	1 (14.3)	0 (0.0)	0 (0.0)	0 (0.0)	3 (3.1)
E. coli	1 (1.8)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (1.0)
Klebsiella spp.	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	0 (0.0)	0 (0.0)	0 (0.0)	1 (1.0)
P. aeruginosa	22 (40.0)	1 (9.1)	5 (41.7)	0 (0.0)	2 (28.6)	0 (0.0)	2 (66.7)	6 (75.0)	38 (38.8)
P. putida	9 (16.4)	0 (0.0)	1 (8.3)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (12.5)	11(11.2)
Total	55 (56.1)	11 (11.2)	12 (12.2)	1 (1.0)	7 (7.1)	1 (1.0)	3 (3.1)	8 (8.2)	98 (100.0)

Table 4. Distribution of bacteria isolates across different clinical presentations of external ocular and periocular infections in a Ghanaian ophthalmic population. *n* frequency, % percentage frequency, *CONS* coagulase-negative staphylococcal species.

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United States of America²⁶ where Gram negatives were found to be significantly lower compared to Gram positive bacteria and with proportional estimates of ranging from six to ten percent^{4,11,13,16,26}. Conversely, the proportionate distribution of Gram negatives to positive bacteria found in this study are parallel with results from several existing literatures^{2,5,8,10}. For example, among the various ocular microbiology investigations conducted across Ethiopia, by Ayebubizu et al.⁵, Belyhun et al.², Assefa et al.¹⁰ as well as Getahun and colleagues⁸ the proportionate

		Bivariate regression			
Variable	OR	95% CI	p-value		
Demographic characteristics		10000	P tutte		
Age (years)	1.04	0.97-1.11	0.237		
Age group (years)					
0-2	Ref				
3-17			0.998		
18-39	-		0.999		
≥40	1.00		1.000		
Gender					
Man	Ref				
Woman			0.998		
Not applicable*			0.998		
Sex designated at birth					
Male	Ref		1		
Female	0.32	0.03-2.97	0.316		
Ethnicity			1		
Akan	Ref				
Northerner			0.999		
Religion					
Catholic	Ref				
Protestant	itei		0.999		
Muslim	1.00		1.000		
Atheist	1.00		1.000		
Socioeconomic characteristics	1.00		1.000		
Residence					
Rural	D.f		1		
	0.99	0.16.6.21	0.002		
Urban	0.99	0.16-6.21	0.993		
Highest level of education		1	1		
None	Ref		1.000		
Preschool	1.00		1.000		
Primary	_		0.999		
Secondary	1.00		0.999		
Tertiary	1.00		1.000		
Not applicable*	1.00		1.000		
Occupation		1	1		
Not applicable*	Ref				
Student	_		0.999		
Trading	1.00		1.000		
Welding	1.00		1.000		
Hairdressing	1.00		1.000		
Unemployed	1.00		1.000		
Construction worker	1.00		1.000		
Beautician	1.00		1.000		
Wage or salary worker	1.00		1.000		
Driver/Transport business	1.00		1.000		
Businessman/woman	1.00		1.000		
Farming	1.00		1.000		
Haidressing	1.00		1.000		
Marital status					
Not applicable*	Ref				
Single	2.24	0.24-21.29	0.481		
Married			0.998		
Caladida			0.999		
Cohabiting	1	1	0.999		
Divorced			0.999		
Divorced			1.000		
•					

	Bivariate regression			
Variable	OR	95% CI	p-value	
Health status variables				
Smoking habits				
Not applicable*	Ref			
Never smoker	6.27	0.67-58.30	0.107	
Past smoker	<u> </u>		0.999	
Current smoker			1.000	
Alcohol intake habits				
Not applicable*	Ref		1	
I never drink	5.67	0.61-52.83	0.128	
I drink only on special occasions	5.07	0.01 52.05	0.999	
I drink once or twice a week			0.999	
			0.999	
Average alcohol consumption per	1		1	
None	Ref		0.000	
1 unit			0.999	
2–5 unit			0.999	
Medical history			-	
Diabetes	1	1	1	
No	Ref			
Yes			0.999	
Hypertension				
No	Ref			
Yes			0.999	
Tuberculosis			·	
No	Ref			
Yes			1.000	
Sexually transmitted diseases				
No	Ref			
Yes			1.000	
Peptic ulcer	1		1	
No	Ref			
Yes			0.999	
Are you currently taking any med	ication			
No	Ref			
Yes			0.998	
Clinical characteristics			0.998	
Eyes affected	D.f		1	
Right eye; oculus dexter	Ref			
Left eye; oculus sinister	0.47	0.04-5.70	0.555	
Both eyes; oculus uterque	1.75	0.15-20.42	0.655	
Site of the eye affected				
Eyelids/lashes		1	1	
No	Ref			
Yes			0.998	
Conjunctiva	_		<u>.</u>	
No	Ref			
Yes			0.999	
Cornea				
No	Ref			
Yes	0.36	0.06-2.31	0.282	
Risk factors		1		
Previous eye infections				
No	Ref			
Yes	0.19	0.02-1.77	0.145	
Previous use of antimicrobials	1		1	
No	Ref	1		
110	Rei		1	

		Bivariate regression		
Variable	OR	95% CI	p-value	
Yes			0.998	
Previous use of contact lens	1			
No	Ref			
Yes			1.000	
Previous use of spectacles				
No	Ref			
Yes	0.46	0.05-4.53	0.505	
Previous ocular trauma	0.40	0.05-4.55	0.505	
No	Ref			
Yes			0.998	
			0.996	
Previous ocular surgery	D C	1	1	
No	Ref			
Yes			0.999	
Duration of illness	1 - 1			
<1 week	Ref		-	
2-4 weeks			0.998	
>4 weeks	1.22	0.13-11.62	0.863	
Previous application of mascara		1		
No	Ref			
Yes			0.999	
Previous application of breastm	ilk			
No	Ref			
Yes			0.999	
Patient presenting symptoms				
Eye pain				
No	Ref			
Yes	0.29	0.03-2.73	0.294	
Itching		1	1	
No	Ref			
Yes	2.88	0.31-26.70	0.352	
Falling lashes	1			
No	Ref			
Yes			1.000	
Lacrimation/watering			1.000	
No	Ref			
Yes	1.10	0.18-6.89	0.920	
Hyperemia/redness	1.10	0.10 0.09	0.920	
No	Ref			
No Yes		0.07.5.04	0.690	
	0.63	0.07-5.84	0.680	
Swelling	n.c		1	
No	Ref		0.000	
Yes			0.998	
Photophobia			1	
No	Ref		-	
Yes	0.32	0.05-2.03	0.315	
Discharge		1		
No	Ref			
Yes	2.27	0.36-14.21	0.381	
Burning sensation				
No	Ref			
Yes			0.999	
Foreign body sensation		ι.		
No	Ref			
Yes				
Others		1	0.999	
			1	

	Bivari regres		
Variable	OR	95% CI	p-value
No	Ref		
Yes	0.56	0.06-5.42	0.615

Table 5. Factors associated with external ocular and periocular infections among a Ghanaian ophthalmic population. Bivariate regression analysis at a significance of p < 0.05. *OR* odds ratio, *CI* confidence interval, *Ref* reference.

distribution of Gram-negatives were similar to Gram positives. Of note, whereas the magnitude of Gram-negative bacteria etiology found in Australia²¹, Iran²⁰ and Italy²⁴ were not equivalent to our findings as well as studies in parts of Ethiopia, the proportion estimates reported were relatively higher. There exist regional variations in the patterns of distribution of Gram-negative bacteria, however, the higher prevalence in our study are ascribed to hygiene as the primary mode of transmission of these enteric bacteria are through oral-fecal contamination. Specifically, we observed during data collection that most patients repeatedly clean ocular discharges with either bare hands or face handkerchief, hence predisposes eyes to contamination by fecal contaminants. Additionally, majority of our study subjects were either in preschool and/or primary hence prone to eye contamination through outdoor gaming activities in school. A considerably higher proportion of the study participants were below two years, and these age categories frequently experience oral-ocular contamination through inserting hands in mouth and touching of eyes thereafter which may have accounted for the increasingly abundance of Gram negative bacteria than positives in our study.

The predominant bacteria species found in our study were *Pseudomonas aeruginosa* and *Staphylococcus aureus*. Although, *S. aureus* was second only to *P. aeruginosa* as the frequently isolated bacteria pathogen, however, it remains the most abundant Gram positive bacteria isolate from all obtainable ocular specimen in our study. This finding are consistent with studies in India¹⁹, Italy²⁴, Nigeria⁵², and Ethiopia^{4,8,12}. The occurrence of ocular infections with *S. aureus* etiology may be due to frequent touching of eyes with filthy hands among study subjects. The incidence of *Pseudomonas aeruginosa* in eye infections are mostly linked with the wearing of contact lens nonetheless we observed an inverse trend in our study. The blinding risk factor associated with *Pseudomonas aeruginosa* ocular infections underscores the promotion of contact lenses as an alternative to spectacle glasses in vision correction and/or cosmesis. Importantly *Pseudomonas aeruginosa* are opportunistic pathogens with devastating consequences on the ocular tissues. Specifically, they induce cornea infiltration and ulcerative keratitis when improperly managed by clinicians. Further, the conjunctiva and cornea are in close proximity landmarked by the limbus, hence pathogens of the conjunctiva can easily spread to the cornea during physiological blinking or mechanical rubbing of the eyes. Given the predominance of *Pseudomonas aeruginosa* in cases of conjunctivitis and keratitis warrants the need for clinicians to probe for other proxy predisposing factors other than relying solely on contact lenses etiology in most instances.

The commonly administered antimicrobial therapy found in this study were Polymyxin B, neomycin and ciprofloxacin. Polymyxin B is a nonribosomal peptidic antimicrobial agents used mostly in the treatment of Gramnegative infections. In particular, they exert their bactericidal effect by binding to phosphate residues within the lipopolysaccharides cell wall to induce displacement of divalent magnesium and calcium cations known to maintain membrane stabilizing properties of Gram-negative bacteria. Consequently, the intrinsic mechanism of action primarily causes an increase in cell membrane permeability resulting in a direct loss of cytoplasmic cell contents. Furthermore, they act synergistically with beta-lactam antibiotics by exposing the peptidoglycan machinery of these Gram negatives for which the latter act on⁵³. On the contrary as an aminoglycoside neomycin actively inhibits protein synthesis of bacteria by insurmountably binding to the 16S ribosomal R.N.A. as well as 50S ribosomal subunits of susceptible class of Gram bacteria^{54,55}. Similarly, ciprofloxacin a fluoroquinolone prevents bacteria D.N.A. replication by terminating the action of the reaction enzymes D.N.A. topoimerase IV and D.N.A. gyrase. The ensued effect is suicidal against Gram negatives as well as mixed bacteria culture⁵⁶. Altogether, the frequent use of the aforementioned antibiotic agents in clinical management of ocular and periocular eye infections in our study are concordant with the laboratory results which identified Gram negatives as the predominant bacteria isolates.

The previous use of antimicrobials among patients is usually considered a risk factor in ocular infection due to the increased potential for contamination from improper handling or storage^{8,57}. Although Getahun et al.⁸ in northwestern Ethiopia reported a significant association between the previous use of antimicrobials and the presence of positive bacterial culture, our results showed otherwise. Thus, patient characteristics, such as the prior usage of antimicrobials, were not significant determinants of positive bacterial culture. This was consistent with a similar investigation by Belyhun and coworkers⁵⁷. Considering the varying resistance mechanism of microorganisms to single antimicrobial therapy and its negative repercussion in resistant eye infections, the administration of two/more specific antibiotic treatment underscores the patronage of a single broad-spectrum antibiotic, an essential determinant in antimicrobial resistance.

Of note, the study has several strengths worth highlighting. The study presents a preliminary and most recent data on bacterial etiology of external ocular and periocular infections among ophthalmic patients in Ghana. Although, we recommend future ocular antibiotic sensitivity studies in this setting, however, in light of the present evidence on the bacterial isolates implicated in eye infection are essential in assisting ophthalmic clinicians in their choice of antibiotic therapy. Moreover, unlike previous studies^{2,7,58} the present investigation

utilized sample from multiple sites which underscores the selection bias usually associated with the convenience sampling approach which the study employed. On the contrary, owing to resource limitation the study could not performed direct fluorescent antibody test and/or Giemsa staining to investigate infections of *Chlamydia trachomatis* etiology. Our prevalence may have been underestimated as a considerable number of our patients where preschoolers whose uncooperative nature denied researchers from taking ocular swabs for bacteriological analyses. Although, given the nature of the studies we could not ascertain the treatment outcomes of the patients the information on the therapy was pivotal in our laboratory.

Conclusion

The prevalence of positive bacteria culture from external ocular and periocular infections was approximately 95%. Gram-negative organisms were commonly implicated and with *Pseudomonas aeruginosa* and *Staphylococcus aureus* as the predominant causative bacteria. Clinical presentations of conjunctivitis and keratitis infections were mostly caused by *Pseudomonas aeruginosa* and with polymyxin B, neomycin and ciprofloxacin as the frequently administered antimicrobial therapy. Given the high burden of ocular bacterial infections, measures (infections control program and antimicrobial agent management program) should be institutionalized to prevent emergence of resistant strains. We recommend future studies to focus on investigating into the potential antibiotic resistances infections within the Ghanaian ophthalmic population.

Methods

Study design, setting and population. A multi-center study was conducted among patients suspected of external ocular and periocular infections in three health facilities in Ghana, namely Anglican Eye Hospital, Jachie; St. Michaels Hospital, Pramso; and Kumasi South Hospital, Agogo from July 18, 2021 to September 18, 2021. Cornea scrapings and conjunctival specimens were obtained from infected eyes for bacterial isolation together with collation of patients sociodemographic and clinical characteristics with a pretested structured questionnaire.

Study setting. The Anglican Eye Hospital (A.E.H.), St. Michaels Hospital (S.M.H.) and Kumasi South Hospital (KSH) were selected for the study primarily because of their higher out-patient-department (O.P.D.) turnout, as well as their wide catchment area and rural/urban interactions. The A.E.H. and S.M.H. are located in Bosomtwe; a rural district in Ghana whilst KSH is situated in the Asokwa municipal area, an urban settlement in the Ashanti Region of Ghana. All the facilities have either a permanent/visiting ophthalmologist, optometrists, ophthalmic nurses and opticians. All the facilities provide comprehensive eye services which range from case history, visual acuity assessment, refraction, dispensing of refractive glasses, management of anterior and posterior segment pathologies, prescribing of medications and performing scheduled surgeries. The S.M.H. and KSH, serve as immediate referral hospitals for the surrounding private and polyclinics. However, all clinical emergencies are referred to the Komfo Anokye Teaching Hospital, the only tertiary health facility in the region. The study facilities lack microbiology laboratory hence clinicians employ empirical approaches in their routine diagnosis and management of external ocular and periocular infections.

Study population and sampling. The study population involved patients who sought ophthalmic treatments/ services for external ocular and periocular infections at the eye clinics of the Anglican Eye Hospital, Jachie; St Michaels Hospital, Pramso; and the Kumasi South Hospital from July 18 to September 18, 2021. Purposive sampling approach was used to recruit all patients presenting with signs and symptoms of external ocular and periocular infections following a consent (and accent for minors). Patients reporting solely for optical correction, or participants on systemic/topical antibiotics or have performed ocular surgery in the last one week were excluded. A purposive sampling approach was used to recruit all one hundred and fourteen (114) eligible subjects.

Study variables. The independent variables for this study were participants' sociodemographic factors; age, sex, ethnicity, religion, facility, socioeconomic; residence, highest level of education, occupation, marital status, health status; smoking habits, alcohol intake, systemic medical conditions and clinical characteristics (eyes affected, site of the eyes affected, risk factors, patient presenting symptoms) whereas the outcome/dependent variable was prevalence of bacteria ocular and periocular infections. Participants were assessed and clinical presentations classified based on operational terms reported previously⁷, and antimicrobial treatments documented accordingly.

Operational definitions. Blepharitis in the study was characterized by gritty itchy sore eyes, with crusting and/or collarattes around the base of the eyelashes coupled with clogging of the Meibomian gland, loss of eyelashes and with demodex conjunctivitis. Conjunctivitis was defined as conjunctival lesion delineated by hyperemia, chemosis, whitish tint purulent discharge and hemorrhage. Blepharoconjuntivitis presented as redness of the eye, with dry scaly eyelids and ensuing symptoms of itchiness and burning sensation. In keratoconjuntivitis, cornea and conjunctiva were implicated with complaints of dryness, itching and mucous discharge. Keratitis was defined as lesion of the cornea with characteristic cornea edema, cellular infiltration, pain, redness, photophobia and ciliary injection. Hordeolum was defined swelling and tenderness of the eyelid with acute pain, photophobia and mild epiphora. Ophthalmia neonatorum is a neonatal conjunctivitis presented within the first 28 days of life with signs of eyelid edema, erythema and purulent discharge. Preseptal and orbital cellulitis showed simi-

lar features of painful swelling and/or tenderness of the eyelid with the later distinguished from the former by decreased vision and pain on eye motility.

Data collection. Sociodemographic and clinical data. The patients sociodemographic, socioeconomic, and health status variables were gathered by the principal investigator and a trained research assistant using a pretested structured questionnaire. A comprehensive vision assessment including visual acuity, slit lamp biomicroscopy, and ophthalmoscopy was performed by a registered optometrist on all study participants. Subsequently, patient medical history, primary and secondary diagnosis, and antimicrobial therapy prescribed were extracted using a data collection form.

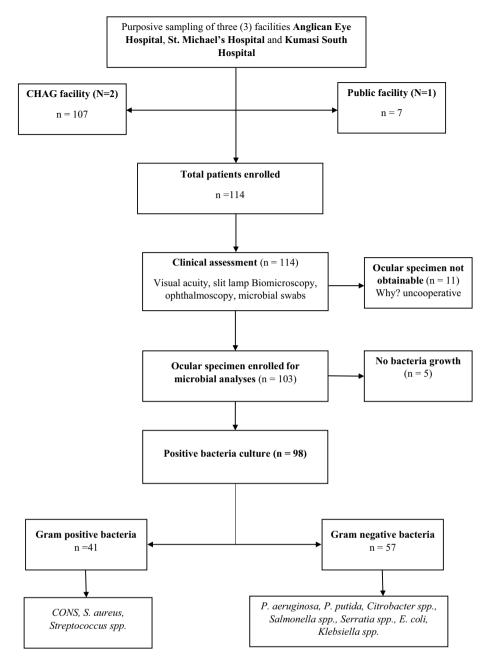
Specimen collection and transport. Overall, 103 ocular specimens were obtained from the eyes of patients with external ocular and periocular infections following aseptic procedures. With the patient eyes in an upward position of gaze, conjunctival specimens were obtained by gently rolling a moistened saline cotton bud over the lower tarsal plate of the eyelids and fornix of the conjunctiva in repeated strokes, thus from nasal to temporal and vice versa. Samples from corneal ulcer and keratitis infections were obtained utilizing a modified version of an original protocol described previously⁸. Briefly, using slit-lamp biomicroscopy and under topical local anesthesia (1–2 drops of 0.5% fresh proparacaine), the edges of the ulcer were firmly scraped^{8,9,19}. On the contrary, none of the patients in our study presented with dacryocystitis and/or blepharitis; hence a puncture and/or aspiration of the lacrimal sac as well as swabbing on the eyelids were not undertaken. The swabs were subsequently kept in a sterile, freshly prepared nutrient broth and transported within 1–3 h in a standard triple packaging system (of an absorbent cotton wrapping primary container enclosed in a sealed bag and kept in an insulated icebox with icepacks) from the study sites to the Microbiology Laboratory of the Faculty of Pharmacy and Pharmaceutical Sciences, Kwame Nkrumah University of Science and Technology, for further microbial investigation.

Laboratory methods. *Culture and identification of bacteria pathogens.* All specimen obtained were initially inoculated on a Nutrient agar and incubated at 37 °C for 24 h and plates examined for growth afterwards. Plates with microbial growth were transferred onto various differential and selective media for preliminary isolation and identification. Specifically, bacteria were cultured on a Mannitol Agar, MacConkey Agar, Bismuth sulphite (all from Oxoid Ltd. Basingstoke, Hants United Kingdom brand), Cetrimide agar (HiMedia Laboratories Pvt. Ltd Mumbai, India), 5% sheep Blood and Chocolate agars and incubated for 24 h. Of note, with the exception of 5% blood and chocolate agars which were kept under anaerobic conditions, an aerobic atmosphere was maintained for all other differential agar media used. Conversely, plates with no growth were re-incubated for additional 48 h and consequently counted negative if no growth pattern appeared. In addition, microbial growth on differential media was again sub-cultured on a nutrient agar to obtain pure colonies and subsequently subjected to further phenotypic identification, specifically colony morphology, Gram stain, and biochemical analyses. Gram-positive bacteria isolates were characterized using coagulase, catalase, and bacitracin tests, whereas Gram-negative bacteria were differentiated using citrate utilization, lysine decarboxylase agar, indole, and urease tests, as well as triple sugar iron agar^{31,59,60}. Details of the methodology are summarized in Fig. 1.

Quality assurance and control. The questionnaires employed to gather the sociodemographic and clinical data were pretested at the Anglican Eye Hospital and revised accordingly following their feedback. Questionnaires were written in English and administered by the principal investigator and a trained research assistant. On the one hand, questionnaire was explained in local dialect for study subjects who could not comprehend instructions in English language. The data from study was doubled checked for accuracy and completeness. At the facility, all test tubes with ocular samples were well-labeled to avoid any mismatched. The laboratory reagents and culture media for the experiments were checked for expiry dates, and sterility control performed to ascertain the integrity of the media such as free from contamination. The media performance and/or functionality assessment was conducted using the American Type Culture Collection (A.T.C.C.) Standard Reference Strains. Specifically, *Escherichia. coli* ATCC 25,922; *Pseudomonas aeruginosa* ATCC 4853; *Staphylococcus aureus* ATCC 25,923.

Data protection and management. The hardcopy version of the filled questionnaires was kept under lock and key and accessible only to the principal investigator and research advisor. Similarly, the softcopy of the non-aggregated dataset was protected using an alphanumeric stringed password. The research report presented to the respective facilities and for publication purposes were aggregated hence individual study participants could not be traced.

Ethical consideration. To undertake this study a hierarchical ethical consideration protocols were followed. A written permission was sought from the authorities of the Anglican Eye Clinic, St. Michaels Hospital, and Kumasi South Hospital. The study protocol was then approved by the Committee on Human Research, Publication and Ethics (C.H.R.P.E.), of the Kwame Nkrumah University of Science and Technology and the Komfo Anokye Teaching Hospital (Reference number: CHRPE/AP/282/21). Written informed consent was obtained from adult participants and for minors a written informed consent was taken from caregivers after study protocol was fully explained to the best of their comprehension. The study adhered to the tenets of the declaration of Helsinki⁶¹, and all laboratory procedures performed in accordance to the Clinical Laboratory Standard Institute guidelines, C.L.S.I.⁶².





Statistical analysis. Data were entered and managed in Microsoft Excel and further exported into Statistical Package and Service Solution version (I.B.M. Corporation IBM* SPSS* Statistics for Windows, version 25.0 Armonk, NY) compatible with windows. Normality assessment was performed using the Kolmogorov Smirnov statistic. The demographics, socioeconomics, health status and clinical characteristics of the sample were presented, and the difference between males and females was tested with chi-square analysis. Clinical diagnosis, cultural status and antimicrobial treatments were presented in cross tabulations using frequencies and percentages. Association between sample characteristics and prevalence of bacterial infections were investigated using bivariate logistic regression at a significance set at p < 0.05 (Supplementary Information S1).

Data availability

All relevant data and materials supporting the conclusion of this article is/are available within the manuscript and its supporting information files.

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References

- Prakash, M. V., Sivakumar, S., Dayal, A., Chitra, A. & Subramaniam, S. Ocular morbidity patterns among children in schools for the blind in Chennai. *Indian J. Ophthalmol.* 65, 733–737. https://doi.org/10.4103/ijo.IJO_294_17 (2017).
- Belyhun, Y. et al. Ocular bacterial infections and antibiotic resistance patterns in patients attending Gondar Teaching Hospital Northwest Ethiopia. BMC Res. Notes 11, 597. https://doi.org/10.1186/s13104-018-3705-y (2018).
- Boadi-Kusi, S. B. et al. A study of Ophthalmia Neonatorum in the Central Reion of Ghana: Causative Agents and Antibiotic Susceptibility Patterns. Glob. Pediatr. Health 8, 2333794X211019700. https://doi.org/10.1177/2333794X211019700 (2021).
- Mshangila, B. et al. External ocular surface bacterial isolates and their antimicrobial susceptibility patterns among pre-operative cataract patients at Mulago National Hospital in Kampala Uganda. BMC Ophthalmol. 13, 71. https://doi.org/10.1186/1471-2415-13-71 (2013).
- Ayehubizu, Z., Mulu, W. & Biadglegne, F. Common bacterial causes of external ocular infections, associated risk factors and antibiotic resistance among patients at ophthalmology unit of Felege Hiwot Referral Hospital, Northwest Ethiopia: A cross-sectional study. J. Ophthal. Inflamm. Infect 11, 7. https://doi.org/10.1186/s12348-021-00238-2 (2021).
- Diriba, K., Kassa, T., Alemu, Y. & Bekele, S. In vitro biofilm formation and antibiotic susceptibility patterns of bacteria from suspected external eye infected patients attending ophthalmology clinic Southwest Ethiopia. *Int. J. Microbiol.* 2020, 8472395. https://doi.org/10.1155/2020/8472395 (2020).
- Mohammed, A. A., Ali, M. M. & Zenebe, M. H. Bacterial etiology of ocular and periocular infections, antimicrobial susceptibility profile and associated factors among patients attending eye unit of Shashemene comprehensive specialized hospital, Shashemene Ethiopia. *BMC Ophthalmol.* 20, 124. https://doi.org/10.1186/s12886-020-01398-w (2020).
- Getahun, E., Gelaw, B., Assefa, A., Assefa, Y. & Amsalu, A. Bacterial pathogens associated with external ocular infections alongside eminent proportion of multidrug resistant isolates at the University of Gondar Hospital, northwest Ethiopia. *BMC Ophthalmol.* 17, 151. https://doi.org/10.1186/s12886-017-0548-6 (2017).
- Teweldemedhin, M., Saravanan, M., Gebreyesus, A. & Gebreegziabiher, D. Ocular bacterial infections at Quiha Ophthalmic Hospital, Northern Ethiopia: An evaluation according to the risk factors and the antimicrobial susceptibility of bacterial isolates. BMC Infect. Dis. 17, 207. https://doi.org/10.1186/s12879-017-2304-1 (2017).
- Assefa, Y. *et al.* Bacteriological profile and drug susceptibility patterns in dacryocystitis patients attending Gondar University Teaching Hospital Northwest Ethiopia. *BMC Ophthalmol.* 15, 34. https://doi.org/10.1186/s12886-015-0016-0 (2015).
- Shiferaw, B., Gelaw, B., Assefa, A., Assefa, Y. & Addis, Z. Bacterial isolates and their antimicrobial susceptibility pattern among patients with external ocular infections at Borumeda hospital Northeast Ethiopia. *BMC Ophthalmol.* 15, 103. https://doi.org/10. 1186/s12886-015-0078-z (2015).
- Muluye, D., Wondimeneh, Y., Moges, F., Nega, T. & Ferede, G. Types and drug susceptibility patterns of bacterial isolates from eye discharge samples at Gondar University Hospital Northwest Ethiopia. *BMC Res. Notes* 7, 292. https://doi.org/10.1186/1756-0500-7-292 (2014).
- Ke, R. et al. Bacteriological profiles and drug susceptibility of Streptococcus isolated from conjunctival sac of healthy children. BMC Pediatr. 20, 306. https://doi.org/10.1186/s12887-020-02203-9 (2020).
- Cho, C. H. & Lee, S. B. Comparison of clinical characteristics and antibiotic susceptibility between Pseudomonas aeruginosa and P. putida keratitis at a tertiary referral center: A retrospective study. BMC Ophthalmol. 18, 204. https://doi.org/10.1186/s12886-018-0882-3 (2018).
- Suwal, S., Bhandari, D., Thapa, P., Shrestha, M. K. & Amatya, J. Microbiological profile of corneal ulcer cases diagnosed in a tertiary care ophthalmological institute in Nepal. BMC Ophthalmol. 16, 209. https://doi.org/10.1186/s12886-016-0388-9 (2016).
- Al-Dhaheri, H. S., Al-Tamimi, M. D., Khandekar, R. B., Khan, M. & Stone, D. U. Ocular pathogens and antibiotic sensitivity in bacterial keratitis isolates at King Khaled Eye Specialist Hospital, 2011 to 2014. Cornea 35, 789–794. https://doi.org/10.1097/ICO. 000000000000844 (2016).
- Wang, N. et al. Bacterial spectrum and antibiotic resistance patterns of ocular infection: differences between external and intraocular diseases. J. Ophthalmol. 2015, 813979. https://doi.org/10.1155/2015/813979 (2015).
- Long, C. et al. Causative organisms of post-traumatic endophthalmitis: A 20-year retrospective study. BMC Ophthalmol. 14, 34. https://doi.org/10.1186/1471-2415-14-34 (2014).
- Bharathi, M. J., Ramakrishnan, R., Shivakumar, C., Meenakshi, R. & Lionalraj, D. Etiology and antibacterial susceptibility pattern of community-acquired bacterial ocular infections in a tertiary eye care hospital in south India. *Indian J. Ophthalmol.* 58, 497–507. https://doi.org/10.4103/0301-4738.71678 (2010).
- Khosravi, A. D., Mehdinejad, M. & Heidari, M. Bacteriological findings in patients with ocular infection and antibiotic susceptibility patterns of isolated pathogens. *Singapore Med. J.* 48, 741–743 (2007).
- Cabrera-Aguas, M., Khoo, P., George, C. R. R., Lahra, M. M. & Watson, S. L. Antimicrobial resistance trends in bacterial keratitis over 5 years in Sydney Australia. *Clin. Exp. Ophthalmol.* 48, 183–191. https://doi.org/10.1111/ceo.13672 (2020).
- 22. Ly, C. N. *et al.* Bacteria commonly isolated from keratitis specimens retain antibiotic susceptibility to fluoroquinolones and gentamicin plus cephalothin. *Clin. Exp. Ophthalmol.* **34**, 44–50. https://doi.org/10.1111/j.1442-9071.2006.01143.x (2006).
- Lee, A. E., Niruttan, K., Rawson, T. M. & Moore, L. S. P. Antibacterial resistance in ophthalmic infections: A multi-centre analysis across UK care settings. BMC Infect. Dis. 19, 768. https://doi.org/10.1186/s12879-019-4418-0 (2019).
- Grandi, G. et al. Bacterial etiology and antimicrobial resistance trends in ocular infections: A 30-year study, Turin area Italy. Eur. J. Ophthalmol. 31, 405–414. https://doi.org/10.1177/1120672119896419 (2021).
- Papa, V., Blanco, A. R. & Santocono, M. Ocular flora and their antibiotic susceptibility in patients having cataract surgery in Italy. J. Cataract. Refract. Surg. 42, 1312–1317. https://doi.org/10.1016/j.jcrs.2016.07.022 (2016).
- Ta, C. N. et al. Antibiotic resistance patterns of ocular bacterial flora: A prospective study of patients undergoing anterior segment surgery. Ophthalmology 110, 1946–1951. https://doi.org/10.1016/s0161-6420(03)00735-8 (2003).
- Teweldemedhin, M., Gebreyesus, H., Atsbaha, A. H., Asgedom, S. W. & Saravanan, M. Bacterial profile of ocular infections: A systematic review. *BMC Ophthalmol.* 17, 212. https://doi.org/10.1186/s12886-017-0612-2 (2017).
- Kowalski, R. P. et al. The prevalence of bacteria, fungi, viruses, and acanthamoeba from 3,004 cases of keratitis, endophthalmitis, and conjunctivitis. Eye Contact Lens 46, 265–268. https://doi.org/10.1097/ICL.00000000000642 (2020).
- Song, X., Xu, L., Sun, S., Zhao, J. & Xie, L. Pediatric microbial keratitis: A tertiary hospital study. Eur. J. Ophthalmol. 22, 136–141. https://doi.org/10.5301/EJO.2011.8338 (2012).
- Bertino, J. S. Jr. Impact of antibiotic resistance in the management of ocular infections: The role of current and future antibiotics. *Clin. Ophthalmol.* 3, 507–521. https://doi.org/10.2147/opth.s5778 (2009).
- 31. Sharma, S. Diagnosis of infectious diseases of the eye. *Eye* (*Lond*) **26**, 177–184. https://doi.org/10.1038/eye.2011.275 (2012).
- Austin, A. *et al.* Empirical treatment of bacterial keratitis: An international survey of corneal specialists. *BMJ Open Ophthalmol.* 2, 1. https://doi.org/10.1136/bmjophth-2016-000047 (2017).
- Machowska, A. & Stalsby Lundborg, C. Drivers of irrational use of antibiotics in Europe. Int. J. Environ. Res. Public Health 16, 1. https://doi.org/10.3390/ijerph16010027 (2018).
- Asbell, P. A. et al. Ocular TRUST: Nationwide antimicrobial susceptibility patterns in ocular isolates. Am. J. Ophthalmol. 145, 951–958. https://doi.org/10.1016/j.ajo.2008.01.025 (2008).

- Haas, W., Pillar, C. M., Torres, M., Morris, T. W. & Sahm, D. F. Monitoring antibiotic resistance in ocular microorganisms: Results from the antibiotic resistance monitoring in ocular microrganisms (ARMOR) 2009 surveillance study. Am. J. Ophthalmol. 152, 567–574. https://doi.org/10.1016/j.ajo.2011.03.010 (2011).
- Metz-Gercek, S. & Mittermayer, H. The European surveillance activities EARSS and ESAC in the context of ABS International. Wien Klin Wochenschr 120, 264–267. https://doi.org/10.1007/s00508-008-0967-8 (2008).
- Molstad, S. et al. Sustained reduction of antibiotic use and low bacterial resistance: 10-year follow-up of the Swedish Strama programme. Lancet Infect. Dis. 8, 125–132. https://doi.org/10.1016/S1473-3099(08)70017-3 (2008).
- Yevutsey, S. K. et al. Situational analysis of antibiotic use and resistance in Ghana: Policy and regulation. BMC Public Health 17, 896. https://doi.org/10.1186/s12889-017-4910-7 (2017).
- Hawkey, P. M. The growing burden of antimicrobial resistance. J. Antimicrob. Chemother. 62(Suppl 1), i1-9. https://doi.org/10. 1093/jac/dkn241 (2008).
- Mainous, A. G., Diaz, V. A., Matheson, E. M., Gregorie, S. H. & Hueston, W. J. Trends in hospitalizations with antibiotic-resistant infections: US, 1997–2006. *Public Health Rep.* 126, 354–360. https://doi.org/10.1177/003335491112600309 (2011).
- 41. Cosgrove, S. E. The relationship between antimicrobial resistance and patient outcomes: Mortality, length of hospital stay, and health care costs. *Clin. Infect. Dis.* **42**(Suppl 2), S82-89. https://doi.org/10.1086/499406 (2006).
- Opintan, J. A. et al. Laboratory-based nationwide surveillance of antimicrobial resistance in Ghana. Infect. Drug Resist. 8, 379–389. https://doi.org/10.2147/IDR.588725 (2015).
- Ding, P., Li, X., Jia, Z. & Lu, Z. Multidrug-resistant tuberculosis (MDR-TB) disease burden in China: A systematic review and spatio-temporal analysis. BMC Infect. Dis. 17, 57. https://doi.org/10.1186/s12879-016-2151-5 (2017).
- Li, B. Y. et al. Rising challenge of multidrug-resistant tuberculosis in China: A predictive study using Markov modeling. Infect. Dis. Poverty 9, 65. https://doi.org/10.1186/s40249-020-00682-7 (2020).
- Barchitta, M. et al. Antibiotic consumption and resistance during a 3-Year Period in Sicily, Southern Italy. Int. J. Environ. Res. Public Health 16, 1. https://doi.org/10.3390/ijerph16132253 (2019).
- JL, O. N. Antimicrobial Resistance: Tackling a crisis for the health and wealth of nations. https://amr-review.org/sites/default/files/ AMR%20Review%20Paper%20-%20Tackling%20a%20crisis%20for%20the%20health%20and%20wealth%20of%20nations_1.pdf. (2014).
- Mohammed, J., Hounmanou, Y. M. G. & Thomsen, L. E. Antimicrobial resistance among clinically relevant bacterial isolates in Accra: A retrospective study. *BMC Res. Notes* 11, 254. https://doi.org/10.1186/s13104-018-3377-7 (2018).
- Kyei, S., Appiah, E., Ayerakwa, E. A., Antwi, C. B. & Asiedu, K. Microbial safety implications of in-use topical diagnostic ophthalmic medications in eye clinics in Ghana. J. Optom. 12, 263–271. https://doi.org/10.1016/j.optom.2019.02.002 (2019).
- Opintan, J. A. & Newman, M. J. Prevalence of antimicrobial resistant pathogens from blood cultures: Results from a laboratory based nationwide surveillance in Ghana. *Antimicrob. Resist. Infect. Control* 6, 64. https://doi.org/10.1186/s13756-017-0221-0 (2017).
- Tesfaye, T. et al. Bacterial profile and antimicrobial susceptibility pattern of external ocular infections in Jimma University specialized hospital. Southwest Ethiopia. 1, 13–20 (2013).
- Shahaby, A. F. & El Tarras, A. Potential bacterial pathogens of red eye infections and their antibiotic susceptibility patterns in Taif. KSA. Int. J. Curr. Microbiol. Appl. Sci. 11, 383–393. https://doi.org/10.2166/wh.2021.274 (2015).
- Okesola, A. O. & Salako, A. O. Microbiological profile of bacterial conjunctivitis in ibadan Nigeria. Ann Ib Postgrad. Med. 8, 20–24. https://doi.org/10.4314/aipm.v8i1.63953 (2010).
- Trimble, M. J., Mlynarcik, P., Kolar, M. & Hancock, R. E. Polymyxin: Alternative mechanisms of action and resistance. Cold Spring Harb. Perspect. Med. 6, 1. https://doi.org/10.1101/cshperspect.a025288 (2016).
- Jana, S. & Deb, J. K. Molecular understanding of aminoglycoside action and resistance. *Appl. Microbiol. Biotechnol.* 70, 140–150. https://doi.org/10.1007/s00253-005-0279-0 (2006).
- Mingeot-Leclercq, M. P., Glupczynski, Y. & Tulkens, P. M. Aminoglycosides: Activity and resistance. Antimicrob. Agents Chemother. 43, 727–737. https://doi.org/10.1128/AAC.43.4.727 (1999).
- Campoli-Richards, D. M. et al. Ciprofloxacin. A review of its antibacterial activity, pharmacokinetic properties and therapeutic use. Drugs 35, 373–447. https://doi.org/10.2165/00003495-198835040-00003 (1988).
- Belyhun, Y. et al. Ocular bacterial infections and antibiotic resistance patterns in patients attending Gondar Teaching Hospital. Northwest Ethiopia. 11, 1–7 (2018).
- Nash, S. D. et al. Ocular Chlamydia trachomatis infection under the surgery, antibiotics, facial cleanliness, and environmental improvement strategy in Amhara, Ethiopia, 2011–2015. Clin. Infect. Dis. 67, 1840–1846. https://doi.org/10.1093/cid/ciy377 (2018).
- 59. Hemavathi, S. Profile of microbial isolates in ophthalmic infections and antibiotic susceptibility of the bacterial isolates: a study in an eye care hospital, bangalore. *J. Clin. Diagn. Res.* **8**, 23–25. https://doi.org/10.7860/JCDR/2014/6852.3910 (2014).
- Hesje, C. K., Sanfilippo, C. M., Haas, W. & Morris, T. W. Molecular epidemiology of methicillin-resistant and methicillin-susceptible Staphylococcus aureus isolated from the eye. *Curr. Eye Res.* 36, 94–102. https://doi.org/10.3109/02713683.2010.534229 (2011).
- 61. Association, W. M. Declaration of Helsinki—ethical principles for medical research involving human subjects. (2013).
- Institute, C. A. L. S. Performance Standards for Antimicrobial Susceptibility Testing; Twenty-Seven Informational Supplement; CLSI Document M02-A12 and M07-A10, CLSI. (2017).

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Author contributions

I.O.D.J: conceptualization; experimental design; project administration; data curation; visualization; methodology; investigation; software; formal analysis; writing original draft; writing—review and editing. T.M.P; project administration; resources; supervision, writing- reviews and editing. L.S.B; project administration; resources; conceptualization; methodology; supervision; writing original draft; writing—review and editing. K.A; project administration; resources; writing—review and editing. F.K.D.A; project administration; resources; methodology; writing—review and editing. D.B.K; resources; writing—review and editing, L.A.O; resources; writing—review and editing. A.D.A; resources; writing—review and editing. B.O.P; methodology; resources; writing—review and editing. S.Y.G; resources; writing—review and editing. J.M; resources; writing—review and editing. C.A.D; resources; writing—review and editing. K.O.A; methodology; data curation; resources; writing original draft; writing reviews and editing. All authors have read and approved the manuscript.

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Competing interests

The authors declare no competing interests.

Additional information

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